5

CLAIMS

What is claimed is:

- 1. A method for raising NO:hemoglobin in red blood cells in a mammal, comprising administering to the mammal a therapeutically effective amount of a composition comprising NO, a composition comprising an inorganic nitrite, a composition comprising an organic nitrite, or a composition comprising a donor of biologically active NO.
- A method for alleviating the effects of a medical intervention in a human, said intervention resulting in a low NO:hemoglobin ratio in red blood cells, said
 method comprising administering to the mammal a therapeutically effective amount of a composition comprising NO, a composition comprising an inorganic nitrite, a composition comprising an organic nitrite, or a composition comprising a donor of biologically active NO.
- The method of Claim 2 in which the medical intervention comprises treatment
 with erythropoietin, treatment by blood transfusion, or treatment with one or more hemoglobin-based blood substitutes.
- A method for treating a condition in a mammal associated with a low NO:hemoglobin ratio in red blood cells, said method comprising administering to the mammal a therapeutically effective amount of a composition comprising NO, a composition comprising an inorganic nitrite, a composition comprising an organic nitrite, or a composition comprising a donor of biologically active NO.

- 5. The method of Claim 4 in which the condition is systemic hypertension, pulmonary hypertension, stroke, myocardial infarction, sickle cell disease, sepsis, thalassemia, polycythemia, a congenital disorder of red blood cells, coronary disease, or a hypoxic condition.
- A method for treating pulmonary hypertension in a mammal, comprising administering to the mammal a therapeutically effective amount of a composition comprising NO, a composition comprising an inorganic nitrite, a composition comprising an organic nitrite, or a composition comprising a donor of biologically active NO.
- A method for treating sickle cell disease in a human, comprising administering to the human a therapeutically effective amount of a composition comprising NO, a composition comprising an inorganic nitrite, a composition comprising an organic nitrite, or a composition comprising a donor of biologically active NO.
- 8. A method for increasing coronary blood flow in a mammal, comprising
 administering to the human a therapeutically effective amount of a composition
 comprising NO, a composition comprising an inorganic nitrite, a composition
 comprising an organic nitrite, or a composition comprising a donor of
 biologically active NO.
- A method for treating a hypoxic condition in a mammal, comprising
 administering to the human a therapeutically effective amount of a composition comprising NO, a composition comprising an inorganic nitrite, a composition comprising an organic nitrite, or a composition comprising a donor of biologically active NO.

15

- 10. A method for raising NO:hemoglobin in red blood cells in a mammal, comprising administering to the mammal a therapeutically effective amount of a composition comprising red blood cells enriched with biologically active NO.
- 11. A method for alleviating the effects of a medical intervention in a human, said medical intervention resulting in a low NO:hemoglobin ratio in red blood cells, said method comprising administering to the human a therapeutically effective amount of a composition comprising red blood cells enriched with biologically active NO.
- The method of Claim 11 in which the medical intervention comprises treatment
 with erythropoietin, treatment by blood transfusion, or treatment with one or
 more hemoglobin-based blood substitutes.
 - 13. A method for treating a condition in a mammal associated with a low NO:hemoglobin ratio in red blood cells, said method comprising administering to the human a therapeutically effective amount of a composition comprising red blood cells enriched with biologically active NO.
 - 14. The method of Claim 13 in which the condition is systemic hypertension, pulmonary hypertension, stroke, myocardial infarction, sickle cell disease, sepsis, thalassemia, polycythemia, a congenital disorder of red blood cells, coronary disease, or a hypoxic condition.
- 20 15. A method for treating pulmonary hypertension in a mammal, comprising administering to the mammal a therapeutically effective amount of a composition comprising red blood cells enriched with biologically active NO.

- 16. A method for treating sickle cell disease in a human, comprising administering to the human a therapeutically effective amount of a composition comprising red blood cells enriched with biologically active NO.
- A method for increasing coronary blood flow in a mammal, comprising
 administering to the mammal a therapeutically effective amount of a
 composition comprising red blood cells enriched with biologically active NO.
 - 18. A method for treating a hypoxic condition in a mammal, comprising administering to the mammal a therapeutically effective amount of a composition comprising red blood cells enriched with biologically active NO.
- 10 19. A method for increasing biologically active NO in isolated red blood cells, comprising contacting the isolated red blood cells with a composition comprising a reagent selected from the group consisting of: NO, an S-nitrosothiol, an ester of an S-nitrosothiol, and ethyl nitrite.
 - 20. The method of Claim 19 wherein the S-nitrosothiol is S-nitrosoglutathione.
- 15 21. A method for potentiating NO-mediated bioactivity of red blood cells, said method comprising administering to the mammal a therapeutically effective amount of a composition comprising a thiol.
 - 22. The method of Claim 21 wherein the thiol is glutathione.
- 23. A method for restoring NO:hemoglobin to a value in a desirable range, in blood for transfusion, said method comprising contacting the blood with a solution

comprising a reagent selected from the group consisting of: NO, an S-nitrosothiol, an ester of an S-nitrosothiol, and ethyl nitrite.

- 24. A method for potentiating the activity of blood for transfusion, comprising adding to the blood a composition comprising one or more thiols.
- A method for identifying a human candidate for a condition associated with a low NO:Hb ratio in red blood cells, said method comprising measuring NO content as S-nitrosothiol and as iron nitrosylhemoglobin in red blood cells isolated from a human, measuring hemoglobin content in the red blood cells isolated from the human, determining a NO:Hb ratio, and comparing the NO:Hb ratio to a normal mean value for NO:Hb, wherein if the NO:Hb ratio is significantly below the normal mean value, the human is a candidate for a condition associated with a low NO:Hb ratio in red blood cells.
- A method for raising NO:hemoglobin as SNO in red blood cells in a mammal to a desirable value, comprising administering to the mammal a therapeutically effective amount of a composition comprising NO, a composition comprising an inorganic nitrite, a composition comprising an organic nitrite, or a composition comprising a donor of biologically active NO.